

## Hit List

|               |                     |       |          |           |
|---------------|---------------------|-------|----------|-----------|
| Clear         | Generate Collection | Print | Fwd Refs | Bkwd Refs |
| Generate OACS |                     |       |          |           |

Search Results - Record(s) 1 through 10 of 12 returned.

☐ 1. Document ID: US 6217851 B1

L4: Entry 1 of 12

File: USPT

Apr 17, 2001

US-PAT-NO: 6217851

DOCUMENT-IDENTIFIER: US 6217851 B1

TITLE: Anti-caries oral compositions

DATE-ISSUED: April 17, 2001

INVENTOR-INFORMATION:

| NAME               | CITY           | STATE | ZIP CODE | COUNTRY |
|--------------------|----------------|-------|----------|---------|
| Kleinberg; Israel  | Smithtown      | NY    |          |         |
| Acevedo; Ana Maria | Caracas        |       |          | VE      |
| Chatterjee; Robi   | South Setanket | NY    |          |         |

US-CL-CURRENT: 424/49; 424/687

|      |       |          |       |        |                |      |           |           |             |        |      |        |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|--------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KWIC | Draw D |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|--------|

☐ 2. Document ID: US 5013542 A

L4: Entry 2 of 12

File: USPT

May 7, 1991

US-PAT-NO: 5013542

DOCUMENT-IDENTIFIER: US 5013542 A

**\*\* See image for Certificate of Correction \*\***

TITLE: Method to inhibit adhesion of disease-causing microorganisms to teeth

DATE-ISSUED: May 7, 1991

INVENTOR-INFORMATION:

| NAME               | CITY    | STATE | ZIP CODE | COUNTRY |
|--------------------|---------|-------|----------|---------|
| Hay; Donald I.     | Wayland | MA    |          |         |
| Gibbons; Ronald J. | Boston  | MA    |          |         |
| Moreno; Edgard G.  | Nahant  | MA    |          |         |

US-CL-CURRENT: 424/54; 514/12, 514/21

|      |       |          |       |        |                |      |           |           |             |        |      |        |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|--------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KWIC | Draw D |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|--------|

☐ 3. Document ID: US 4683292 A

L4: Entry 3 of 12

File: USPT

Jul 28, 1987

US-PAT-NO: 4683292

DOCUMENT-IDENTIFIER: US 4683292 A

TITLE: Immunotherapeutic polypeptide agents which bind to lymphocyte immunoglobulin FC receptors

DATE-ISSUED: July 28, 1987

## INVENTOR-INFORMATION:

| NAME          | CITY      | STATE | ZIP CODE | COUNTRY |
|---------------|-----------|-------|----------|---------|
| Hahn, Gary S. | San Diego | CA    |          |         |

US-CL-CURRENT: 530/328; 930/10, 930/20, 930/DIG.785, 930/DIG.788, 930/DIG.802, 930/DIG.811

|      |       |          |       |        |                |      |           |           |             |        |      |         |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|---------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KWIC | Draw. D |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|---------|

☐ 4. Document ID: US 4585757 A

L4: Entry 4 of 12

File: USPT

Apr 29, 1986

US-PAT-NO: 4585757

DOCUMENT-IDENTIFIER: US 4585757 A

**\*\* See image for Certificate of Correction \*\***

TITLE: Hypotensive active peptides

DATE-ISSUED: April 29, 1986

## INVENTOR-INFORMATION:

| NAME                   | CITY    | STATE | ZIP CODE | COUNTRY |
|------------------------|---------|-------|----------|---------|
| Pang, Peter K. T.      | Lubbock | TX    |          |         |
| Tenner, Jr.; Thomas E. | Lubbock | TX    |          |         |

US-CL-CURRENT: 514/18; 514/19, 930/10, 930/20, 930/70, 930/DIG.820

|      |       |          |       |        |                |      |           |           |             |        |      |         |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|---------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KWIC | Draw. D |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|---------|

☐ 5. Document ID: US 4499068 A

L4: Entry 5 of 12

File: USPT

Feb 12, 1985

US-PAT-NO: 4499068

DOCUMENT-IDENTIFIER: US 4499068 A

**\*\* See image for Certificate of Correction \*\***

TITLE: Oral compositions comprising N.sup.G -alkyl derivatives of arginine

DATE-ISSUED: February 12, 1985

## INVENTOR-INFORMATION:

| NAME                 | CITY       | STATE | ZIP CODE | COUNTRY |
|----------------------|------------|-------|----------|---------|
| Silbering; Steven B. | Plainsboro | NJ    |          |         |
| Sipos; Tibor         | Lebanon    | NJ    |          |         |

US-CL-CURRENT: 424/52; 424/54, 562/560

|      |       |          |       |        |                |      |           |           |             |        |      |          |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KMOC | Draw. De |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|

☐ 6. Document ID: US 4499067 A

L4: Entry 6 of 12

File: USPT

Feb 12, 1985

US-PAT-NO: 4499067

DOCUMENT-IDENTIFIER: US 4499067 A

TITLE: Oral compositions comprising N.sup.G -acyl derivatives of arginine

DATE-ISSUED: February 12, 1985

## INVENTOR-INFORMATION:

| NAME                 | CITY       | STATE | ZIP CODE | COUNTRY |
|----------------------|------------|-------|----------|---------|
| Silbering; Steven B. | Plainsboro | NJ    |          |         |
| Sipos; Tibor         | Lebanon    | NJ    |          |         |

US-CL-CURRENT: 424/52; 424/54, 554/38, 554/47, 554/53, 562/560

|      |       |          |       |        |                |      |           |           |             |        |      |          |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KMOC | Draw. De |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|

☐ 7. Document ID: US 4486403 A

L4: Entry 7 of 12

File: USPT

Dec 4, 1984

US-PAT-NO: 4486403

DOCUMENT-IDENTIFIER: US 4486403 A

TITLE: Composition for and treatment of teeth

DATE-ISSUED: December 4, 1984

## INVENTOR-INFORMATION:

| NAME              | CITY        | STATE | ZIP CODE | COUNTRY |
|-------------------|-------------|-------|----------|---------|
| Mechanic; Gerald  | Chapel Hill | NC    | 27514    |         |
| Binderman; Itzhak | Tel-Aviv    |       |          | IL      |

US-CL-CURRENT: 424/54; 424/49

|      |       |          |       |        |                |      |           |           |             |        |      |          |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KMOC | Draw. De |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|

☐ 8. Document ID: US 4477429 A

L4: Entry 8 of 12

File: USPT

Oct 16, 1984

US-PAT-NO: 4477429

DOCUMENT-IDENTIFIER: US 4477429 A

TITLE: Oral compositions comprising N.sup..alpha. -alkyl derivatives of arginine

DATE-ISSUED: October 16, 1984

## INVENTOR-INFORMATION:

| NAME                 | CITY       | STATE | ZIP CODE | COUNTRY |
|----------------------|------------|-------|----------|---------|
| Silbering; Steven B. | Plainsboro | NJ    |          |         |
| Sipos; Tibor         | Lebanon    | NJ    |          |         |

US-CL-CURRENT: 424/52; 424/54, 562/560

|      |       |          |       |        |                |      |           |           |             |        |      |          |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KWMC | Draw. De |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|

☐ 9. Document ID: US 4477428 A

L4: Entry 9 of 12

File: USPT

Oct 16, 1984

US-PAT-NO: 4477428

DOCUMENT-IDENTIFIER: US 4477428 A

TITLE: Oral compositions comprising N.sup..alpha.,N.sup.G -diacyl derivatives of arginine

DATE-ISSUED: October 16, 1984

## INVENTOR-INFORMATION:

| NAME                 | CITY       | STATE | ZIP CODE | COUNTRY |
|----------------------|------------|-------|----------|---------|
| Silbering; Steven B. | Plainsboro | NJ    |          |         |
| Sipos; Tibor         | Lebanon    | NJ    |          |         |

US-CL-CURRENT: 424/52; 424/54, 554/106, 554/107, 562/560

|      |       |          |       |        |                |      |           |           |             |        |      |          |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KWMC | Draw. De |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|----------|

☐ 10. Document ID: US 4339431 A

L4: Entry 10 of 12

File: USPT

Jul 13, 1982

US-PAT-NO: 4339431

DOCUMENT-IDENTIFIER: US 4339431 A

TITLE: Anticalculus oral composition

DATE-ISSUED: July 13, 1982

## INVENTOR - INFORMATION:

| NAME          | CITY     | STATE | ZIP CODE | COUNTRY |
|---------------|----------|-------|----------|---------|
| Gaffar; Abdul | Somerset | NJ    |          |         |

US-CL-CURRENT: 424/54; 424/49

|      |       |          |       |        |                |      |           |           |             |        |      |            |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|------------|
| Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Claims | KWIC | Draw. Data |
|------|-------|----------|-------|--------|----------------|------|-----------|-----------|-------------|--------|------|------------|

Clear

Generate Collection

Print

Fwd Refs

Bkwd Refs

Generate OACS

Terms

Documents

4225579

12

Display Format: CIT

Change Format

[Previous Page](#)[Next Page](#)[Go to Doc#](#)

## Hit List

Clear

Generate Collection

Print

Fwd Refs

Bkwd Refs

Generate OACS

**Search Results - Record(s) 11 through 12 of 12 returned.**☐ 11. Document ID: US 4339430 A

L4: Entry 11 of 12

File: USPT

Jul 13, 1982

US-PAT-NO: 4339430

DOCUMENT-IDENTIFIER: US 4339430 A

TITLE: Antibacterial oral composition

DATE-ISSUED: July 13, 1982

INVENTOR-INFORMATION:

| NAME          | CITY     | STATE | ZIP CODE | COUNTRY |
|---------------|----------|-------|----------|---------|
| Gaffar; Abdul | Somerset | NJ    |          |         |

US-CL-CURRENT: 424/54; 424/49

Full

Title

Citation

Front

Review

Classification

Date

Reference

Sequences

Attachments

Claims

KWC

Draw. D

☐ 12. Document ID: US 4225579 A

L4: Entry 12 of 12

File: USPT

Sep 30, 1980

US-PAT-NO: 4225579DOCUMENT-IDENTIFIER: US 4225579 A

TITLE: Means and method for improving defenses against caries

DATE-ISSUED: September 30, 1980

INVENTOR-INFORMATION:

| NAME              | CITY      | STATE | ZIP CODE | COUNTRY |
|-------------------|-----------|-------|----------|---------|
| Kleinberg; Israel | Smithtown | NY    | 11787    |         |

US-CL-CURRENT: 424/48; 260/1, 424/54, 514/2, 530/330, 530/331, 930/10

Full

Title

Citation

Front

Review

Classification

Date

Reference

Sequences

Attachments

Claims

KWC

Draw. D

Clear

Generate Collection

Print

Fwd Refs

Bkwd Refs

Generate OACS

Terms

Documents

4225579

12

**Display Format:** CIT

Change Format

[Previous Page](#)

[Next Page](#)

[Go to Doc#](#)

First Hit   Fwd Refs

Generate Collection

Print

L4: Entry 3 of 12

File: USPT

Jul 28, 1987

US-PAT-NO: 4683292

DOCUMENT-IDENTIFIER: US 4683292 A

TITLE: Immunotherapeutic polypeptide agents which bind to lymphocyte immunoglobulin FC receptors

DATE-ISSUED: July 28, 1987

## INVENTOR-INFORMATION:

| NAME          | CITY      | STATE | ZIP CODE | COUNTRY |
|---------------|-----------|-------|----------|---------|
| Hahn; Gary S. | San Diego | CA    |          |         |

## ASSIGNEE-INFORMATION:

| NAME             | CITY      | STATE | ZIP CODE | COUNTRY | TYPE CODE |
|------------------|-----------|-------|----------|---------|-----------|
| Immunetech, Inc. | San Diego | CA    |          |         | 02        |

APPL-NO: 06/ 522602   [PALM]

DATE FILED: August 12, 1983

INT-CL: [04] C07K 7/06

US-CL-ISSUED: 530/328

US-CL-CURRENT: 530/328; 930/10, 930/20, 930/DIG.785, 930/DIG.788, 930/DIG.802, 930/DIG.811

FIELD-OF-SEARCH: 260/112.5R

PRIOR-ART-DISCLOSED:

## U.S. PATENT DOCUMENTS

Search Selected

Search ALL

Clear

|                          | PAT-NO         | ISSUE-DATE     | PATENTEE-NAME    | US-CL      |
|--------------------------|----------------|----------------|------------------|------------|
| <input type="checkbox"/> | <u>3778426</u> | December 1973  | Najjar           | 260/112.5R |
| <input type="checkbox"/> | <u>4153688</u> | May 1979       | Dimicoli et al.  | 260/112.5R |
| <input type="checkbox"/> | <u>4161522</u> | July 1979      | Hamburger        | 260/112.5R |
| <input type="checkbox"/> | <u>4171299</u> | October 1979   | Hamburger        | 260/112.5R |
| <input type="checkbox"/> | <u>4201770</u> | May 1980       | Stevens          | 424/177    |
| <input type="checkbox"/> | <u>4215112</u> | July 1980      | Goldstein et al. | 260/112.5R |
| <input type="checkbox"/> | <u>4223016</u> | September 1980 | Roy et al.       | 260/112.5R |
| <input type="checkbox"/> | <u>4225579</u> | September 1980 | Kleinberg        | 260/112.5R |
|                          | <u>4284537</u> | August 1981    | Beachey          | 260/6      |



|                          |                |               |                  |            |
|--------------------------|----------------|---------------|------------------|------------|
| <input type="checkbox"/> |                |               |                  |            |
| <input type="checkbox"/> | <u>4341755</u> | July 1982     | Lindall          | 424/1      |
| <input type="checkbox"/> | <u>4369138</u> | January 1983  | Lindall          | 260/112.5R |
| <input type="checkbox"/> | <u>4388233</u> | June 1983     | Bissell et al.   | 548/159    |
| <input type="checkbox"/> | <u>4407948</u> | October 1983  | Goodman et al.   | 435/91     |
| <input type="checkbox"/> | <u>4409141</u> | October 1983  | Noda et al.      | 260/112.5R |
| <input type="checkbox"/> | <u>4409144</u> | October 1983  | Heinicke         | 260/112.5R |
| <input type="checkbox"/> | <u>4415493</u> | November 1983 | Weigle et al.    | 260/112.5R |
| <input type="checkbox"/> | <u>4436874</u> | March 1984    | Aspisi et al.    | 525/327.1  |
| <input type="checkbox"/> | <u>4454121</u> | June 1984     | Beachey          | 260/112.5R |
| <input type="checkbox"/> | <u>4457867</u> | July 1984     | Ishida           | 260/112.5R |
| <input type="checkbox"/> | <u>4474757</u> | October 1984  | Arnon et al.     | 424/88     |
| <input type="checkbox"/> | <u>4476116</u> | October 1984  | Anik             | 260/112.5R |
| <input type="checkbox"/> | <u>4497801</u> | February 1985 | Hashimoto et al. | 260/112.5R |

## OTHER PUBLICATIONS

Proc. Nat. Acad. Sci., vol. 72, No. 6, pp. 2081-2083 (1975).  
 Molecular Immunology, vol. 19, No. 10, pp. 1245-1254 (1982).  
 Science, (1975) pp. 389-390, vol. 189.  
 Veretennikova et al., Int. J. Peptide Protein Res. 17, (1981) 430-435.  
 Plummer, et al., Fed. Proc. 42 713 (1983).  
 Hunninghake et al. Clin. Immunol Rev. 1 (3) 337 (1981-1982).  
 Hunsicker, J. Exp. Med. 150 413 (1979).  
 Johnson et al., (J. Immunol., 117, 1491 (1975)).  
 Boackle et al., (Nature, 282, 742 (1979)).  
 Prystowsky et al. (Biochemistry 20, 6349 (1981)).  
 Lukas et al. (J. Immunol. 127, 2555 (1981)).  
 Burton et al. Nature 288, 338 (1980).  
 Morgan et al. (Proc. Natl. Acad. Sci. USA 79, 5388 (1982)).  
 Ciccimarra et al. (Proc. Natl. Acad. Sci. USA 72 208 (1975)).  
 Stanworth Mol. Immunol. 19 1245 (1982).  
 Hamburger, Immunology 38, 78 (1979).  
 Biochem, J., 180 665 (1979).  
 Biochem, J., 181 623 (1979).  
 Barnett-Foster et al. Mol. Immunol 19 407 (1982).  
 Barnett-Foster et al. J. Immunol 120, 407 (1978).  
 Takatsu et al., J. Immunol 114 1838 (1975).  
 Dorrington et al. Immunol Rev. 41, 3 (1978).  
 Perex-Montfort Mol. Immunol 19 1113 (1982).  
 Kuehl et al. Science 210 978 (1980).  
 Goodwin Clin, Immunol Immunopath 15 106 (1980).  
 Stenson et al. Immunol 125 1 (1980).  
 Leung et al. J. Immunol 129 1742 (1982).  
 Fischer et al. J. Immunol 126 1452 (1981).  
 Klein et al. Immunol 48 337 (1983).  
 Goodwin et al. Cancer Immunol Immunother., 8, 3 (1980).  
 Samuelsson Science 220 568 (1983).  
 Weiss et al. J. Immunol 129 309 (1982).  
 Fantone, J. Pathol. 107 397 (1982).  
 Perex et al. Textbook Rheumatology, vol. 1, W. B. Saunders Philadelphia, 1981 pp. 179-194).  
 Dreisn et al. N. Engl. J. Med. 298 358 (1978).

Lawrence et al. N. Engl. J. Med. 302 1187 (1980).  
Holdsworth J. Immunol 130 735 (1983).  
Striker J. Exp. Med. 149, 127 (1979).  
Kumar in Pathologic Basis of Disease eds. S. L. Robbins & R. S. Cotran (W. B. Saunders; Philadelphia, 1979), p. 304.  
Melwicz et al., Clin. Exp. Immunol., 49, 364 (1982).  
Spiegelberg, et al., 42, 124 (1983).  
Scott et al., Fed. Proc., 42, 129 (1983).  
McMillan, N. Engl. J. Med., 304, 1135 (1981).  
Fehr et al. (N. Engl. J. Med., 306, 1254 (1982)).  
Imbach et al. (Lancet, Jun. 6, 1981, p. 1228).  
Oberbarnscheidt et al., Immunol., 35, 151 (1978).  
Kolsch et al., Immunol. Rev., 49, 61 (1980).  
Fridman et al., Immunol Rev. 56, 51 (1981).  
Bich-Thuy, J. Immunol., 129, 150 (1982).  
Smolen et al., J. Immunol., 129, 10150 (1982).  
Goeken et al. Hum. Immunol., 6, 79 (1983).  
Kabelitz et al., Eur. J. Immunol., 12, 687 (1982).  
Sakane et al., Proc. Natl. Acad. Sci. U.S.A., 75, 3464 (1978).  
Miyasaka et al., J. Clin. Invest., 66, 928 (1980).  
James et al., J. Clin. Invest., 66, 1305 (1980).  
Hodgkin's Lymphoma (Engleman, et al., J. Clin. Invest., 66 149 (1980).  
Smith et al. J. Natl. Cancer Inst., 58, 579 (1977).  
Cochrane et al., Lancet 1, 441 (1976).  
Douvas, Ann. Immunol. Inst. Pasteur, 132C, 307 (1981).  
Ulcerative colitis (Hibi, et al., Clin. Exp. Immunol. 49, 75 (1982)).  
Hashimoto's thyroiditis (Calder et al., Clin. Exp. Immunol., 14, 153 (1973)).  
Gonzalez-Molina et al., J. Clin. Invest., 59, 616 (1977).  
Merrifield J. Am. Chem Soc., 85, 2149-2154 (1963).  
Barany & Merrifield in The Peptides eds. E. Gross & F. Meinehofer, vol. 2 (Academic Press, 1980) pp. 2 285.  
Synthesis of a Tetrapeptide by R. B. Merrifield, Journal of American Chemical Society (vol. 85, pp. 85, pp. 2 2154 (1963)).

ART-UNIT: 153

PRIMARY-EXAMINER: Phillips; Delbert R.

ATTY-AGENT-FIRM: Lyon & Lyon

ABSTRACT:

An active site peptide which blocks immune complex binding to Fc receptors, the peptide having an amino acid sequence selected from the group consisting of:

A-B-C-D-E-F-G-H-I-J-K-L-M-N-O-P,

or a subgroup thereof,

wherein

A is Arg, Lys, Orn, Gln, or His;

B is Ser, Thr, Ala, or Gly;

C is Thr, Ser, Ala, or Gly;

D is Thr, Ser, Ala, or Gly;

E is Lys, Arg, Orn or His;

F is Thr, Ser, Ala, or Gly;

G is Ser, Thr, Ala, or Gly;

H is Gly, Ala, Thr, Ser, Lys, Arg, or Orn

I is Pro, Val, Leu, Ile, or Ala;

J is Arg, Lys, Orn, or His;

K is Ala, Thr, Ser, or Gly;

L is Ala, Thr, Ser, or Gly;

M is Pro, Val, Leu, Ile, or Ala;

N is Glu, or Asp;

O is Val, Leu, Ile, or Ala;

P is Tyr, or Phe.

and pharmaceutically acceptable salts thereof.

1 Claims, 4 Drawing figures

[First Hit](#)   [Fwd Refs](#)

Generate Collection

Print

L3: Entry 42 of 57

File: USPT

Nov 11, 1997

US-PAT-NO: 5686075

DOCUMENT-IDENTIFIER: US 5686075 A

TITLE: Synthetic peptide vaccines for dental caries

DATE-ISSUED: November 11, 1997

## INVENTOR-INFORMATION:

| NAME               | CITY        | STATE | ZIP CODE | COUNTRY |
|--------------------|-------------|-------|----------|---------|
| Taubman; Martin A. | Newtonville | MA    |          |         |
| Smith; Daniel J.   | Natick      | MA    |          |         |

US-CL-CURRENT: 424/197.11; 424/185.1, 424/190.1, 424/193.1, 424/194.1, 530/324,  
530/350

## CLAIMS:

We claim:

1. An immunogenic composition comprising a peptide consisting of at least one amino acid sequence selected from the group consisting of:

a) DGKLRYDANS GDQAFN KSV (SEQ ID NO: 4), and

b) PLDKRSGLNPLIHNSLVDREVDDRE (SEQ ID NO: 2); and a physiologically compatible carrier.

2. An immunogenic composition comprising at least two peptides, wherein at least one peptide consists of an amino acid sequence of either DANFDSIRVDAVDNVDADLLQI (SEQ ID NO: 1) or PLDKRSGLNPLIHNSLVDREVDDRE (SEQ ID NO: 2) where both sequences are of the catalytic domain of streptococcal glucosyltransferase, and at least one peptide consists of an amino acid sequence of either TGAQTIKGQKLYFKANGQQVK (SEQ ID NO: 3) or DGKLRYDANS GDQAFN KSV (SEQ ID NO: 4) where both sequences are of the glucan-binding domain of streptococcal glucosyltransferase, and a physiologically compatible carrier.

3. An immunogenic composition of claim 2 where 2 or more of said peptides of the streptococcal glucosyltransferase protein are present and attached to a core matrix of 3 or more lysines.

4. The immunogenic composition of claim 1 wherein said composition induces in a mammal an immune response that is both a B cell response and a T cell response.

5. The immunogenic composition of claim 4 wherein the B cell immune response produces antibodies of the IgG or the IgA isotype.

6. An immunogenic composition comprising at least two peptides of a streptococcal

glucosyltransferase protein covalently attached to a lysine core matrix, wherein each peptide consists of an amino acid sequence selected from the group consisting of:

- a) DANFDSIRVDAVDNVDADLLQI (SEQ ID NO: 1),
- b) TGAQTIKGQKLYFKANGQQVKG (SEQ ID NO: 3),
- c) DGKLRYDANSBGDQAFNKS (SEQ ID NO: 4),
- d) QWNGESEKPYDDHL (SEQ ID NO: 5), and
- e) PLDKRSGLNPLIHNSLVDREVDRE (SEQ ID NO: 2); and

a physiologically compatible carrier.

7. The immunogenic composition of claim 6 having at least one additional immunologic component, which produces an immunogenic response against an infectious organism, covalently attached to said lysine core matrix, wherein said additional immunogenic component is a peptide comprising an amino acid sequence from an immunologic domain selected from the group consisting of diphtheria, pertussis, tetanus and measles.

8. The immunogenic composition of claim 6 wherein the lysine core matrix consists of at least three lysines.

9. The immunogenic composition of claim 6 wherein said composition induces in a mammal an immune response that is a B cell response, a T cell response or both a B cell response and a T cell response.

10. The immunogenic composition of claim 9 wherein both the B cell response and T cell response are elicited by the same amino acid sequence.

11. The immunogenic composition of claim 10 wherein the B cell immune response produces antibodies of the IgG or the IgA isotype.

12. An immunogenic composition of claim 6 comprising 4 peptides, where

a) the 4 peptides are the same or different;

b) each peptide consists of an amino acid sequence selected from the group consisting of DANFDSIRVDAVDNVDADLLQI (SEQ ID NO: 1), PLDKRSGLNPLIHNSLVDREVDRE (SEQ ID NO: 2) where both sequences are of the catalytic domain of streptococcal glucosyltransferase, the amino acid sequence DGKLRYDANSBGDQAFNKS (SEQ ID NO: 4) of the glucan binding domain of streptococcal glucosyltransferase, and the amino acid sequence QWNGESEKPYDDHL (SEQ ID NO: 5) of the native streptococcal glucosyltransferase surface domain; and

c) the 4 peptides are attached to a core matrix of 3 lysines.

13. An immunogenic composition of claim 12 wherein said composition induces in a mammal an immune response that results in the reduction of the colonization or accumulation of mutants streptococcal strains in a mammal to whom the immunogenic composition is administered.

14. An immunogenic composition comprising a peptide consisting of an amino acid sequence of PLDKRSGLNPLIHNSLVDREVDDRE (SEQ ID NO: 2) and a physiologically compatible carrier.

15. A method of interfering with the enzymatic activity of streptococcal glucosyltransferase in a mammal comprising the administration of a peptide consisting of an amino acid sequence of PLDKRSGLNPLIHNSLVDREVDDRE (SEQ ID NO: 2) to a mammal in a manner that raises an immune response in the mammal, thereby interfering with the enzymatic activity of streptococcal glucosyltransferase in the mammal.

16. A method of provoking an immune response to streptococcal glucosyltransferase in a mammal comprising the administration of a peptide consisting of an amino acid sequence of either PLDKRSGLNPLIHNSLVDREVDDRE (SEQ ID NO: 2) or DGKLRYDANSQDQAFNKS (SEQ ID NO: 4) in a manner that raises an immune response in the mammal.

17. The method of claim 16 wherein said immune response results in reduction of the colonization or accumulation of mutans, streptococcal strains in the mammal to whom the peptide is administered.

18. An immunogenic composition comprising a peptide consisting of an amino acid sequence of DGKLRYDANSQDQAFNKS (SEQ ID NO: 4) and a physiologically compatible carrier.

19. A method of interfering with the glucan-binding activity of streptococcal glucosyltransferase in a mammal comprising the administration of a peptide consisting of an amino acid sequence of DGKLRYDANSQDQAFNKS (SEQ ID NO: 4) in a manner that induces a response which thereby interferes with the glucan-binding activity of streptococcal glucosyltransferase in the mammal.

20. An immunogenic composition comprising at least two peptides covalently attached to at least one additional immunologic component which produces an immunogenic response against an infectious organism, wherein each peptide is selected from the group consisting of:

a) DANFDSIRVDAVDNVDADLLQI (SEQ ID NO: 1);

b) PLDKRSGLNPLIHNSLVDREVDDRE (SEQ ID NO: 2);

c) TGAQTIKGQKLYFKANGQQVKG (SEQ ID NO: 3);

d) DGKLRYDANSQDQAFNKS (SEQ ID NO: 4); and

e) QWNGESEKPYDDHL (SEQ ID NO: 5); and

a physiologically compatible carrier, wherein said additional immunologic component is a peptide comprising an amino acid sequence from an immunologic domain selected from the group consisting of diphtheria, pertussis, tetanus and measles.

21. An immunogenic composition comprising a peptide covalently attached to at least one additional immunologic component which produces an immunogenic response against an infectious organism, wherein said peptide is selected from the group consisting of:

- a) DANFDSIRVDAVDNVDADLLQI (SEQ ID NO: 1);
- b) PLDKRSGLNPLIHNSLVDREVDDRE (SEQ ID NO: 2);
- c) TGAQTIKGQKLYFKANGQQVKG (SEQ ID NO: 3);
- d) DGKLRYDANSQDQAFNKS (SEQ ID NO: 4); and
- e) QWNGESEKPYDDHL (SEQ ID NO: 5); and

a physiologically compatible carrier, wherein said additional immunologic component is a peptide comprising an amino acid sequence from an immunologic domain selected from the group consisting of diphtheria, pertussis, tetanus and measles.

[First Hit](#)   [Fwd Refs](#)☐  

L3: Entry 42 of 57

File: USPT

Nov 11, 1997

US-PAT-NO: 5686075

DOCUMENT-IDENTIFIER: US 5686075 A

TITLE: Synthetic peptide vaccines for dental caries

DATE-ISSUED: November 11, 1997

## INVENTOR-INFORMATION:

| NAME               | CITY        | STATE | ZIP CODE | COUNTRY |
|--------------------|-------------|-------|----------|---------|
| Taubman; Martin A. | Newtonville | MA    |          |         |
| Smith; Daniel J.   | Natick      | MA    |          |         |

## ASSIGNEE-INFORMATION:

| NAME                                  | CITY   | STATE | ZIP CODE | COUNTRY | TYPE CODE |
|---------------------------------------|--------|-------|----------|---------|-----------|
| Forsyth Dental Infirmary for Children | Boston | MA    |          |         | 02        |

APPL-NO: 08/ 057162   [PALM]

DATE FILED: April 30, 1993

## PARENT-CASE:

RELATED APPLICATION This application is a continuation-in-part of application Ser. No. 07/877,295, now abandoned, entitled "Synthetic Peptide Vaccines for Dental Caries" by Martin A. Taubman and Daniel J. Smith, filed May 1, 1992. The teachings of application Ser. No. 07/877,295 now abandoned, are incorporated herein by reference.

INT-CL: [06] A61 K 39/09

US-CL-ISSUED: 424/197.11; 124/185.1, 124/190.1, 124/193.1, 124/194.1, 530/324, 530/350

US-CL-CURRENT: 424/197.11; 424/185.1, 424/190.1, 424/193.1, 424/194.1, 530/324, 530/350

FIELD-OF-SEARCH: 424/88, 424/92, 424/185.1, 424/190.1, 424/193.1, 424/194.1, 424/197.11, 424/244.1, 530/350, 530/324, 530/325, 530/326

PRIOR-ART-DISCLOSED:

## U.S. PATENT DOCUMENTS

  

|                          | PAT-NO         | ISSUE-DATE    | PATENTEE-NAME  | US-CL   |
|--------------------------|----------------|---------------|----------------|---------|
| <input type="checkbox"/> | <u>4150116</u> | April 1979    | Taubman et al. | 424/88  |
| <input type="checkbox"/> | <u>4250262</u> | February 1981 | Taubman et al. | 435/193 |
|                          | <u>4438200</u> | March 1984    | Taubman et al. | 435/193 |





4894229

January 1990

Polson et al.

424/92

## FOREIGN PATENT DOCUMENTS

| FOREIGN-PAT-NO | PUBN-DATE   | COUNTRY | US-CL |
|----------------|-------------|---------|-------|
| 0328403        | August 1989 | EP      |       |
| WO91/07979     | June 1991   | WO      |       |

## OTHER PUBLICATIONS

Ferretti et al., "Nucleotide Sequence of a Glucosyltransferase Gene from *Streptococcus sorbrinus* MFe28," *Journal of Bacteriology* 169(9): 4271-4278 (1987).

Banas et al., "Sequence Analysis of the Gene for the Glucan-Binding Protein of *Streptococcus mutans* Ingbritt," *Infection and Immunity*, 58(3): 667-673 (1990).

R.R.B. Russell et al., "Homology of Glucosyltransferase Gene and Protein Sequences from *Streptococcus sobrinus* and *Streptococcus mutans*," *J. Dent. Res.*, 67(3): 543-547 (1988, Mar.).

M.T. Dertzbaugh et al., "Cholera Toxin B-Subunit Fusion: Structural and Functional Analysis of the Chimeric Protein," *Infection and Immunity*, 58(1): 70-79 (1990, Jan.).

Taubman, M. et al., "T Cell Epitopes on Synthetic Peptides from the Glucan Binding and Catalytic Regions of Mutants streptococci", Abstract and poster presentation at IADR General Session, Glasgow, Scotland, Jul. 1-4, 1992 (abstract pub. *J. Dent. Res.* 71, p. 577, abstract #491 (1992)).

Ellis in *Vaccines* Chapter 29 Plutkin et al. Eds WB Saunders Co p. 573, 1988.

Taubman et al *Infection and Immunity* vol. 63 No. 8:3088-3093, Aug. 1995.

Smith et al *Infection and Immunity* vol. 55 No. 5 1274-1278, May 1987.

J.P. Tam, "Synthetic Peptide Vaccine Design: Synthesis and Properties of a High Density Multiple Antigenic Peptide System," *Proc. Nat'l Acad. Sci. USA*, 85: 5409-5413 (1988, Aug.).

T. Lehner et al., "Local Oral Immunization with Synthetic Peptides Induces a Dual Mucosal IgG and Salivary IgA Antibody Response and Prevents Colonization of *Streptococcus mutans*," *Immunology*, 67: 419-424 (1989).

Samith et al., "Immunological Characteristics of Synthetic Peptides Derived from Glucosyltransferase (GTF) Sequences . . ." Abstract and poster presentation for Cariology for the Nineties, Rochester, NY, Jun. 4-7, 1991.

Dertzbaugh et al., "Inhibition of *Streptococcus mutans* Glucosyltransferase Activity by Antiserum to a Subsequence Peptide," *Infection and Immunity*, 58(6): 1509-1513 (1990).

Arnon et al. (1992) *FASEB J.* vol. 6 pp. 3265-3274 "Structural Basis of Antigenic Specificity and Design of New Vaccines".

Smith et al (1993) *Infect Immun* 61(7): 2899-2905.

Abo et al (1991) *J. Bact* 173(3):989-996.

Smith et al (1991, Mar.) Published Abstract IADRI AADR, 1991 Apr.-Acapulco, Mexico.

Mooser et al (1991) *J. Biol. Chem.* 266(14):8916-8922.

Mosci et al (1989) *Minerva Stomatal* 38(3):379-388 (Abstract Only).

Wong et al (1990) *Infect Immun* 58(7):2165-2170.

Honda et al (1990) *J. Gen. Microbiol.* 136:2099-2105.

Shiroza et al (1987) *J. Bact.* 169(9) 4263-4207.

Ueda et al (1988) *Gene* 69: 101-109.

Mooser et al (1988) *Infected Immun* 56(4):880-884.

Hajishengallin et al (1989) *Odontostomatol Proodos* 43(4):315-321 (Abstract Only).

Schneerson et al (1984) *Inf. Immun* 45(3):582-591.

Gregory et al (1987) *Infect. & Immun* 55(10):2409-2415.

Smith et al (1987) *Infect. Immun* 55(11):2562-2569.

ART-UNIT: 186

PRIMARY-EXAMINER: Feisee; Lila

ASSISTANT-EXAMINER: Reeves; Julie E.

ATTY-AGENT-FIRM: Hamilton, Brook, Smith & Reynolds, P.C.

ABSTRACT:

Immunization of animals with a composition containing either an amino acid sequence from the catalytic domain of glucosyltransferase, an amino acid sequence from the glucan-binding region of glucosyltransferase or an amino acid sequence from the native surface domain of glucosyltransferase provoke antibody and T-cell immune responses to this enzyme. Since this enzyme has been implicated in the colonization of mutans streptococci on tooth surfaces, such immune responses are important for the prevention of dental caries. Multicomponent and multivalent compositions which include these amino acid sequences provide effective vaccine capabilities.

21 Claims, 1 Drawing figures

First Hit   Fwd Refs☐ **Generate Collection** **Print**

L3: Entry 50 of 57

File: USPT

May 7, 1991

US-PAT-NO: 5013542

DOCUMENT-IDENTIFIER: US 5013542 A

**\*\* See image for Certificate of Correction \*\***

TITLE: Method to inhibit adhesion of disease-causing microorganisms to teeth

DATE-ISSUED: May 7, 1991

## INVENTOR-INFORMATION:

| NAME               | CITY    | STATE | ZIP CODE | COUNTRY |
|--------------------|---------|-------|----------|---------|
| Hay; Donald I.     | Wayland | MA    |          |         |
| Gibbons; Ronald J. | Boston  | MA    |          |         |
| Moreno; Edgard G.  | Nahant  | MA    |          |         |

US-CL-CURRENT: 424/54; 514/12, 514/21

## CLAIMS:

What is claimed is:

1. A method of inhibiting the adhesion of microorganisms to a mineral surface, which method comprises:

contacting the mineral surface with an adhesive-inhibiting amount of a non-immunogenic, acidic, amino-terminal segment of an anionic proline-rich protein to inhibit the adhesion of disease-causing microorganisms to the mineral surface.

2. The method of claim 1 wherein the mineral surface comprises a manual calcium-containing surface.

3. The method of claim 1 wherein said mineral surface comprises hydroxyapatite.

4. The method of claim 1 wherein the mineral surface comprises a tooth surface.

5. The method of claim 1 wherein the anionic proline-rich protein comprises PRP 1-4, PIF-s or PIF-f.

6. The method of claim 1 which includes cleaving the proline-rich protein by enzymatic cleaving to obtain the acidic amino acid end segment of the protein and contacting the mineral surface with said cleaved amino-terminal segment.

7. The method of claim 1 wherein said segment consists essentially of the first 30 amino-acid residue of the said protein.

8. The method of claim 1 wherein said end segment comprises PCA--ASP--LEU--ASP--GLU--

ASp--VAL--P-Ser--GLN13 GLU--ASP--VAL--PRO--LEU--VAL--ILE--SER--ASP--GLY--GLY--ASP--P-SER--GLU--GL N--PHE--ILE--ASP--GLU--GLU--ARG.

9. The method of claim 1 which includes incorporating the said segment in a pharmaceutically acceptable carrier to form a composition and treating an apatitic surface of a patient with said composition.

10. The method of claim 9 which includes treating the oral cavity of a patient with said composition.

11. The contacted mineral surface prepared by the method of claim 1.

12. A method of inhibiting the adhesion of disease-causing microorganisms, which method comprises:

a) providing as an active ingredient a non-immunogenic peptide consisting essentially of the first acidici 30-residue amino-terminal segment of a proline-rich protein;

b) incorporating the said segment in a pharmaceutically acceptable carrier to form a composition; and

c) introducing the said composition into the oral cavity of a patient.

13. The treated oral cavity prepared by the method of claim 12.

14. The method of claim 1 which includes cleaving the proline-rich protein by enzymatic cleaving by the use of trypsin.

15. The method of claim 1 wherein the disease causing microorganisms are selected from the group consisting of: Streptococcus mutans, Streptococcus sanguis, Streptococcus sobrinus, Actinomyces viscosus and Bacteroides gingivalis.

16. The method of claim 12 wherein the 30 residue amino terminal segment of a proline-rich protein comprises: PCA--ASP--LEU--ASP--GLU--ASP--VAL--P-SER--GLN--GLU--ASP--VAL--PRO--LEU--VA L--ILE--SER--ASP--GLY--GLY--ASP--P-SER--GLU--GLN--PHE--ILE--ASP--GLU--GLU-- ARG.

[First Hit](#)   [Fwd Refs](#)☐ [Generate Collection](#) [Print](#)

L3: Entry 52 of 57

File: USPT

Feb 12, 1985

US-PAT-NO: 4499068

DOCUMENT-IDENTIFIER: US 4499068 A

**\*\* See image for Certificate of Correction \*\***TITLE: Oral compositions comprising N.sup.G -alkyl derivatives of arginine

DATE-ISSUED: February 12, 1985

## INVENTOR-INFORMATION:

| NAME                 | CITY       | STATE | ZIP CODE | COUNTRY |
|----------------------|------------|-------|----------|---------|
| Silbering; Steven B. | Plainsboro | NJ    |          |         |
| Sipos; Tibor         | Lebanon    | NJ    |          |         |

US-CL-CURRENT: 424/52; 424/54, 562/560

## CLAIMS:

We claim:

1. N.sup.G -alkyl derivatives of arginine having the formula: ##STR6## where y is an integer of from 9 to 19, and the pharmaceutically acceptable salts thereof.
2. A compound of claim 1 wherein said pharmaceutically acceptable salts are selected from the group consisting of alkali metal salts, alkaline earth metal salts, amphoteric metal salts, heavy metal salts, organic base salts, and organic and inorganic acid salts.
3. The compound of claim 1 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -decylarginine.
4. The compound of claim 1 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -laurylarginine.
5. The compound of claim 1 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -myristylarginine.
6. The compound of claim 1 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -palmitylarginine.
7. The compound of claim 1 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -stearylarginine.
8. A composition of matter for oral hygiene to inhibit acid production in the oral cavity comprising an effective amount, in a pharmaceutically acceptable carrier, of an N.sup.G -alkyl derivative of arginine having the formula; ##STR7## wherein y is an integer of from 9 to 19, or a pharmaceutically acceptable salt thereof.

9. The composition of matter of claim 8 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -decylarginine.

10. The composition of matter of claim 8 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -laurylarginine.

11. The composition of matter of claim 8 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -myristylarginine.

12. The composition of matter of claim 8 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -palmitylarginine.

13. The composition of matter of claim 8 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -stearylarginine.

14. The composition of matter of claim 8 wherein said pharmaceutically acceptable carrier is a dentifrice.

15. The composition of matter of claim 8 wherein said pharmaceutically acceptable carrier is a lozenge.

16. A composition of matter for oral hygiene to inhibit the formation of caries comprising, in a pharmaceutically acceptable carrier, from about 0.0001% to about 10% of a fluoride salt and an effective amount of an N.sup.G -alkyl derivative of arginine having the formula: ##STR8## wherein y is an integer of from 0 to 29, or a pharmaceutically acceptable salt thereof.

17. The composition of matter of claim 16 wherein said pharmaceutically acceptable carrier is a mouthrinse.

18. The composition of matter of claim 16 wherein said pharmaceutically acceptable carrier is a dentifrice.

19. A composition of matter for oral hygiene to inhibit the formation of caries comprising from about 0.05 to about 10% of N.sup.G -alkyl derivative of arginine having the formula: ##STR9## wherein y is an integer of from 5 to 19, or a pharmaceutically acceptable salt thereof, in combination with from about 0.001 to about 1.0% of a fluoride salt in a pharmaceutically acceptable polyol-containing vehicle.

20. The composition of matter of claim 19 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -laurylarginine.

21. The composition of matter of claim 19 wherein said N.sup.G -alkyl derivative of arginine is N.sup.G -myristylarginine.

22. A method for inhibiting acid production by microorganisms in the oral cavity which comprises introducing into the oral cavity in a pharmaceutically acceptable carrier, an effective amount of an N.sup.G -alkyl derivative of arginine having the formula: ##STR10## wherein y is an integer of from 0 to 29, or a pharmaceutically acceptable salt thereof.

23. A method for inhibiting acid production by microorganisms in the oral cavity which comprises

introducing into the oral cavity a composition comprising, in a pharmaceutically acceptable carrier, from about 0.0001% to about 10% of a fluoride salt and an effective amount of an N.sup.G -alkyl derivative of arginine having the formula: ##STR11## wherein y is an integer of from 0 to 29, or a pharmaceutically acceptable salt thereof.

24. A method for inhibiting acid production by microorganisms in the oral cavity which comprises introducing into the oral cavity a composition comprising from about 0.05 to about 10% of N.sup.G -alkyl derivative of arginine having the formula: ##STR12## wherein y is an integer of from 5 to 19, or a pharmaceutically acceptable salt thereof, in combination with from about 0.001 to about 1.0% of a fluoride salt in a pharmaceutically acceptable polyol-containing vehicle.

[First Hit](#)   [Fwd Refs](#)

Generate Collection

Print

L3: Entry 56 of 57

File: USPT

Mar 15, 1983

US-PAT-NO: RE31181

DOCUMENT-IDENTIFIER: US RE31181 E

TITLE: Means and method for improving natural defenses against caries

DATE-ISSUED: March 15, 1983

## INVENTOR-INFORMATION:

| NAME              | CITY      | STATE | ZIP CODE | COUNTRY |
|-------------------|-----------|-------|----------|---------|
| Kleinberg; Israel | Smithtown | NY    | 11787    |         |

US-CL-CURRENT: 514/18; 260/1, 424/49, 424/52, 424/54, 514/19, 530/330, 530/331

## CLAIMS:

What is claimed is

1. A method for supplementing the body's resistance to caries which comprises providing to the mouth an effective amount of a caries combatting pH rise factor which is .[a peptide.]. .Iadd.a source of pH adjusting compound or precursor thereof .Iaddend.having 2-4 amino acid units at least one of which is arginine.
2. A method as set forth in claim 1 wherein the pH rise factor is provided in concentrations of from about 0.05 mM to about 3 mM.
3. A method as set forth in claim 1 wherein the pH rise factor is provided in combination with a dental care product.
4. A method as set forth in claim 1 wherein the pH rise factor is provided in a food product.
5. A method as set forth in claim 1 wherein the pH rise factor is provided in combination with chewing gum. .Iadd. 6. The method according to claims 1, 3, 4 or 5 wherein said pH-rise factor is applied to the mouth in association with fluoride ions. .Iaddend..Iadd. 7. The method according to claim 1 wherein said pH-rise factor is provided to the mouth in a mouth wash. .Iaddend..Iadd. 8. The method according to claim 1 wherein said pH-rise factor is provided to the mouth in tooth paste. .Iaddend..Iadd. 9. The method according to claim 1 wherein said pH-rise factor is provided in tooth powder..Iaddend.